

5     **Aventis Behring GmbH**

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10    **Abstract:**

**Modified cDNA Factor VIII and its Derivatives**

15    Modified human factor VIII cDNA wherein mutations are inserted either in the wild-type factor VIII cDNA or in a factor VIII cDNA in which the B-domain is partially or completely deleted and may be replaced by a DNA linker segment and

20       A) one or several codons of the human factor VIII cDNA which are not identical with the corresponding codon in the same position of the porcine factor VIII cDNA are substituted by a different codon in such a way that

- 25       •     when the human sequence contains a codon for a neutral amino acid whereas the porcine sequence contains a codon for a charged amino acid then a codon for an amino acid with the same charge as found in the porcine sequence is introduced into the human sequence;
- 30       •     when the human sequence contains a codon for a charged amino acid whereas the porcine sequence contains a codon for a neutral amino acid then a codon for a neutral amino acid or a codon for an amino acid of the opposite charge is introduced into the human sequence,
- 35       •     when the human sequence contains a codon for a charged amino acid whereas the porcine sequence contains a codon for an amino

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acid with the opposite charge then a codon for an amino acid with the opposite charge is introduced into the human sequence or

B) one or several codons for a charged amino acid which are found in the FVIII cDNA of a hemophilic patient are replaced by a codon for an amino acid of the opposite charge.

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